



STIC Search Report

Biotech-Chem Library

STIC Database Tracking Number: 10/723144

TO: Marcela Cordero Garcia
Location: REM/3A30/3C18
Art Unit: 1654
Thursday, May 11, 2006

Case Serial Number: 10/723144

From: Noble Jarrell
Location: Biotech-Chem Library
Rem 1B71
Phone: 272-2556

Noble.jarrell@uspto.gov

Search Notes

Scientific and Technical Information Center
SEARCH REQUEST FORM

Requester's Full Name: MARCELA M CORDERO GARCIA Examiner #: P0381 Date: 5/15/06
 Art Unit: 1654 Phone Number: 2-2939 Serial Number: 10/723,144
 Location (Bldg/Room#): REM3A30 (Mailbox #): 3C18 Results Format Preferred (circle): PAPER DISK

To ensure an efficient and quality search, please attach a copy of the cover sheet, claims, and abstract or fill out the following:

Title of Invention: SEE BIB D5

Inventors (please provide full names): SEE BIB D5

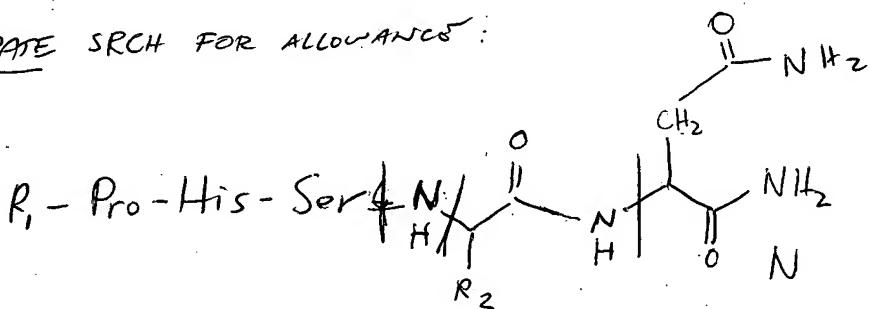
Earliest Priority Date: 11/02

Search Topic:

Please provide a detailed statement of the search topic, and describe as specifically as possible the subject matter to be searched. Include the elected species or structures, keywords, synonyms, acronyms, and registry numbers, and combine with the concept or utility of the invention. Define any terms that may have a special meaning. Give examples or relevant citations, authors, etc., if known.

For Sequence Searches Only Please include all pertinent information (parent, child, divisional, or issued patent numbers) along with the appropriate serial number.

PLEASE UPDATE SRCH FOR ALLOWANCE



R_1 = acyl, substituted acyl, oxy carbonyl & substituted oxy carbonyl

R_2 = alkyl, $-(\text{CH}_2)_m \text{S(O)}_n \text{R}'$ or $-(\text{CH}_2)_m \text{S(O)}_n-\text{S(O)}_n \text{R}'$

$m = 1$ or 2 , $n & \sigma = 0, 1$ or 2

R' = OPEN (can be alkyl, substituted alkyl, acyl, substituted acyl, aryl, substituted aryl, arylalkyl, substituted arylalkyl, heteroalkyl, substituted heteroalkyl, heteroaryl, substituted heteroaryl, heteroarylalkyl, subst. heteroarylalkyl, oxy carbonyl, substituted oxy carbonyl)

***** STAFF USE ONLY *****

Searcher: Noble

Type of Search

Vendors and cost where applicable

Searcher Phone #: _____

NA Sequence (#)

STN Dialog

Searcher Location: _____

AA Sequence (#)

Questel/Orbit Lexis/Nexis

Date Searcher Picked Up: 5/11/06

Structure (#)

Westlaw WWW/Internet

Date Completed: 5/11/06

Bibliographic

In-house sequence systems

'Searcher Prep & Review Time: 15

Litigation

Commercial Oligomer Score/Length
 Interference SPDI Encode/Transl
 Other (specify) _____

Online Time: 25

Fulltext

Other

=> b reg
FILE 'REGISTRY' ENTERED AT 07:01:43 ON 11 MAY 2006
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STRUCTURE FILE UPDATES: 9 MAY 2006 HIGHEST RN 883631-57-0
DICTIONARY FILE UPDATES: 9 MAY 2006 HIGHEST RN 883631-57-0

New CAS Information Use Policies, enter HELP USAGETERMS for details.

TSCA INFORMATION NOW CURRENT THROUGH January 6, 2006

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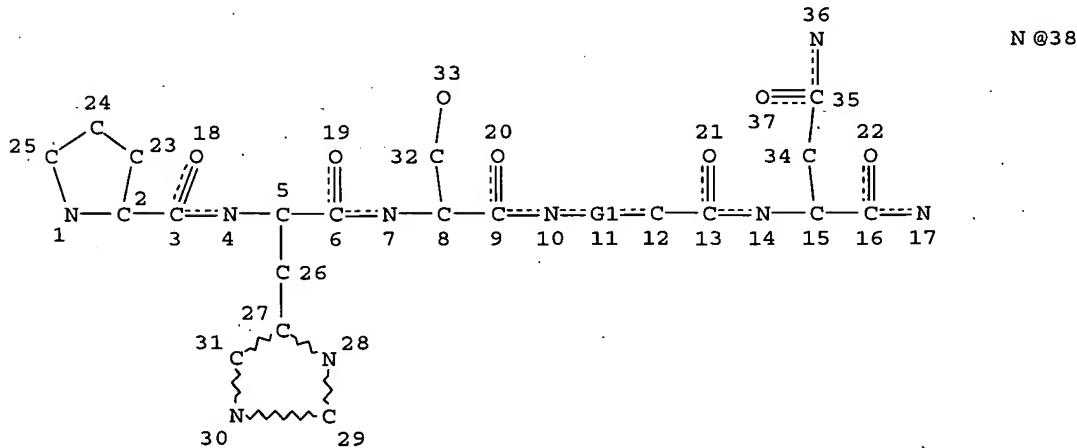
*
* The CA roles and document type information have been removed from *
* the IDE default display format and the ED field has been added, *
* effective March 20, 2005. A new display format, IDERL, is now *
* available and contains the CA role and document type information. *
*

Structure search iteration limits have been increased. See HELP SLIMITS
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on property searching in REGISTRY, refer to:

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=> d que sta 124
L22 STR



REP G1=(0-1) 38
NODE ATTRIBUTES:
CONNECT IS M1 RC AT 1
CONNECT IS M1 RC AT 12
CONNECT IS M1 RC AT 38
DEFAULT MLEVEL IS ATOM

DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:

RSPEC 27

NUMBER OF NODES IS 38

STEREO ATTRIBUTES: NONE

L24 64 SEA FILE=REGISTRY CSS FUL L22

100.0% PROCESSED 33412 ITERATIONS

64 ANSWERS

SEARCH TIME: 00.00.01

=> b hcap

FILE 'HCAPLUS' ENTERED AT 07:01:56 ON 11 MAY 2006

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FILE COVERS 1907 - 11 May 2006 VOL 144 ISS 20

FILE LAST UPDATED: 9 May 2006 (20060509/ED)

New CAS Information Use Policies, enter HELP USAGETERMS for details.

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d all fhitstr l26 tot

L26	ANSWER 1 OF 3	HCAPLUS	COPYRIGHT 2006 ACS on STN		
AN	2004:610128	HCAPLUS			
DN	141:157478				
ED	Entered STN:	30 Jul 2004			
TI	Peptides which target tumor and endothelial cells, compositions and uses thereof				
IN	Allan, Amy L.; Yoon, Won Hyung; Gladstone, Patricia L.; Ternansky, Robert J.; Parry, Graham; Donate, Fernando; Mazar, Andrew				
PA	Attenuon, Llc, USA				
SO	PCT Int. Appl., 117 pp.				
	CODEN: PIXXD2				
DT	Patent				
LA	English				
IC	ICM C07K				
CC	34-3 (Amino Acids, Peptides, and Proteins) Section cross-reference(s): 1, 63				
FAN.CNT	2				
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LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO,
 NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ,
 TM, TN, TR, TT, TZ, UA, UG, UZ, VC, VN, YU, ZA, ZM, ZW
 RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
 BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
 ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK,
 TR; BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
 CA--2506813 AA 20040729 2003CA-2506813 20031125 <--
 US2004162239 A1 20040819 2003US-0723144 20031125 <--
 US2005020810 A1 20050127 2003US-0722843 20031125 <--
 EP---1569678 A2 20050907 2003EP-0796483 20031125 <--
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 BR2003016550 A 20051004 2003BR-0016550 20031125 <--
 CN---1741808 A 20060301 CN 2003-80109204 20031125 <--
 CN---1741809 A 20060301 CN 2003-80109205 20031125 <--
 NO2005003112 A 20050805 2005NO-0003112 20050624 <--
 PRAI 2002US-429174P P 20021125 <--
 2003US-475539P P 20030602 <--
 2003WO-US37895 W 20031125

CLASS

PATENT NO.	CLASS	PATENT FAMILY CLASSIFICATION CODES
WO 2004063213	ICM	C07K
	IPCI	C07K [ICM, 7]
	IPCR	A61K0038-10 [I,A]; A61K0038-10 [I,C]; C07K0007-00 [I,C]; C07K0007-08 [I,A]
CA---2506813	ECLA	C07K007/06A
	IPCI	A61K0038-10 [ICM, 7]; C07K0007-06 [ICS, 7]; A61K0038-08 [ICS, 7]; C07K0007-08 [ICS, 7]; C07K0007-00 [ICS, 7]
	IPCR	A61K0038-10 [I,A]; A61K0038-10 [I,C]; C07K0007-00 [I,C]; C07K0007-08 [I,A]
US2004162239	ECLA	C07K007/06A
	IPCI	A61K0038-08 [ICM, 7]; A61K0038-10 [ICS, 7]; C07K0007-08 [ICS, 7]; C07K0007-06 [ICS, 7]; C07K0007-00 [ICS, 7]
	IPCR	A61K0038-08 [I,A]; A61K0038-08 [I,C]; A61K0038-10 [I,A]; A61K0038-10 [I,C]; C07K0007-00 [I,C]; C07K0007-06 [I,A]; C07K0007-08 [I,A]
US2005020810	NCL	514/012.000
	IPCI	C07K0007-08 [ICM, 7]; C07K0007-06 [ICS, 7]; C07K0007-00 [ICS, 7]
	IPCR	C07K0007-00 [I,C]; C07K0007-06 [I,A]; C07K0007-08 [I,A]
	NCL	530/324.000
EP---1569678	IPCI	A61K0038-10 [ICM, 7]; A61K0038-08 [ICS, 7]; C07K0007-08 [ICS, 7]; C07K0007-06 [ICS, 7]; C07K0007-00 [ICS, 7]
	IPCR	A61K0038-10 [I,A]; A61K0038-10 [I,C]; C07K0007-00 [I,C]; C07K0007-08 [I,A]
BR2003016550	ECLA	C07K007/06A
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	IPCR	A61K0038-10 [I,A]; A61K0038-10 [I,C]; C07K0007-00 [I,C]; C07K0007-08 [I,A]
CN---1741808	ECLA	C07K007/06A
	IPCI	A61K0038-10 [I,A]; A61K0038-08 [I,A]; C07K0007-06 [I,A]; C07K0007-08 [I,A]; C07K0007-00 [I,C]
CN---1741809	IPCI	A61K0038-10 [I,A]; A61K0038-08 [I,A]; C07K0007-08 [I,A]; C07K0007-06 [I,A]; C07K0007-00 [I,C]
NO2005003112	IPCI	A61K0038-04 [ICM, 7]; C07K0007-08 [ICS, 7]; C07K0007-06 [ICS, 7]; C07K0007-00 [ICS, 7]
	ECLA	C07K007/06A

OS MARPAT 141:157478

AB The invention relates generally to peptide analogs of Ac-PHSCN-NH₂ which target tumor and endothelial cells and have antitumor, antiangiogenic and antimetastatic activity and to methods for their synthesis and use in pharmaceutical compns. for treating, preventing and detecting diseases characterized by tumor growth, metastasis and angiogenesis. The peptide

analogs may serve, inter alia, as carriers of radioactivity, PET-active compds., toxins, fluorescent mols. and PEG mols. Peptides R1[(NHCHR2CO)0-1(X1)0-100]m-X2-X3-X4-X5-X6-[(X7)0-1(NHCHR3CO)0-1]nNR4R5 [R1 is (un)substituted acyl, alkyl, cycloalkyl or imino, or acyl chelate; R2 is substituted alkyl; R4, R5 are (un)substituted alkyl; X1, X7 are NH(CH:CH)1-6CO, NH(CH2)1-6CO, NHCHMeCO; X2-X6 are α -amino acids which are defined; m, n are 0 or 1, with the proviso that R1 is not acetyl when R4 and R5 are H and m and n are 0] are claimed. Thus, Ac-Pro-His-Ser-Cys(Ac)-Asn-OH was prepared by the solid-phase method and coupled to doxorubicin hydrochloride to afford the conjugate.

ST peptide prolylhistidylserylcysteinylaspartamide analog prepn antitumor

IT Angiogenesis

Angiogenesis inhibitors

Antitumor agents

Neoplasm

(preparation of peptides which target tumor and endothelial cells)

IT Peptides, preparation

RL: DGN (Diagnostic use); PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of peptides which target tumor and endothelial cells)

IT Polyoxyalkylenes, reactions

RL: RCT (Reactant); RACT (Reactant or reagent)

(preparation of peptides which target tumor and endothelial cells)

IT 729594-60-9P

RL: DGN (Diagnostic use); PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(preparation of peptides which target tumor and endothelial cells)

IT 7440-74-6DP, Indium, complexes with DPTA peptide conjugate

262438-43-7DP, analogs 729594-61-0P 729594-62-1P

729594-63-2P 729594-64-3P 729594-65-4P 729594-66-5P

729594-67-6P 729594-68-7P 729594-69-8P

729594-70-1P 729594-71-2P 729594-72-3P 729594-73-4P

729594-74-5P 729594-75-6P 729594-76-7P 729594-77-8P

729594-78-9P 729594-79-0P 729594-80-3P 729594-81-4P

729594-82-5P 729594-83-6P 729594-84-7P

729594-85-8P 729594-86-9P 729594-87-0P

729594-88-1P 729594-89-2P 729594-90-5P 729594-91-6P

729594-92-7P 729594-93-8P 729594-94-9P

729594-95-0P 729594-96-1P 729594-97-2P

729594-98-3P 729594-99-4P 729595-00-0P 729595-01-1P

729595-02-2P 729595-03-3DP, polyethylene glycol derivative

729595-04-4P 729595-05-5P 729595-06-6P 729595-07-7P

729595-08-8P 729595-09-9P 729595-14-6P 730960-54-0P

731003-01-3DP, Indium complexes 731003-01-3P

731003-02-4P

RL: DGN (Diagnostic use); PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of peptides which target tumor and endothelial cells)

IT 456-22-4, 4 Fluorobenzoic acid 501-97-3 553-12-8 3301-79-9, 6

Carboxyfluorescein 13811-11-5 25316-40-9, Doxorubicin hydrochloride

34071-95-9 66134-67-6 76823-03-5, 5 Carboxyfluorescein 106966-68-1

137076-54-1, 1,4,7,10-Tetraazacyclododecane-1,4,7,10-tetraacetic acid,

tris 1 1 dimethylethyl ester 517913-89-2 622405-78-1 729595-15-7

729595-16-8D, resin-bound 729595-17-9

RL: RCT (Reactant); RACT (Reactant or reagent)

(preparation of peptides which target tumor and endothelial cells)

IT 729595-10-2DP, resin-bound 729595-11-3DP, resin-bound 729595-12-4DP,

resin-bound 729595-13-5DP, resin-bound

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of peptides which target tumor and endothelial cells)

IT 262438-43-7DP, analogs

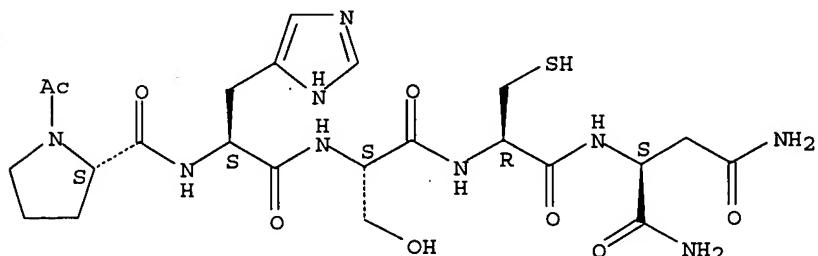
RL: DGN (Diagnostic use); PAC (Pharmacological activity); SPN (Synthetic

preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
 (preparation of peptides which target tumor and endothelial cells)

RN 262438-43-7 HCPLUS

CN L-Aspartamide, 1-acetyl-L-prolyl-L-histidyl-L-seryl-L-cysteinyl- (9CI)
 (CA INDEX NAME)

Absolute stereochemistry.



L26 ANSWER 2 OF 3 HCPLUS COPYRIGHT 2006 ACS on STN

AN 2004:467702 HCPLUS

DN 141:33798

ED Entered STN: 10 Jun 2004

TI Peptides which inhibit angiogenesis, cell migration, cell invasion and cell proliferation, their preparation, and compositions and therapeutic uses thereof

IN Allan, Amy L.; Donate, Fernando; Hopkins, Stephanie A.; Gladstone, Patricia L.; Mazar, Andrew ; O'Hare, Sean M.; Parry, Graham; Plunkett, Marian L.; Ternansky, Robert J.; Yoon, Won Hyung

PA Attenuon, LLC, USA

SO PCT Int. Appl., 88 pp.

CODEN: PIXXD2

DT Patent

LA English

IC ICM A61K

CC 1-8 (Pharmacology)

Section cross-reference(s): 34, 63

FAN.CNT 2

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
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PI WO2004047771	A2	20040610	2003WO-US38175	20031125 <--
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US2004162239	A1	20040819	2003US-0723144	20031125 <--
US2005020810	A1	20050127	2003US-0722843	20031125 <--
BR2003016523	A	20051018	2003BR-0016523	20031125 <--
EP---1594521	A2	20051116	2003EP-0812058	20031125 <--
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JP2006514116	T2	20060427	2005JP-0510345	20031125 <--

NO2005003111	A	20050824	2005NO-0003111	20050624 <--
PRAI 2002US-429174P	P	20021125	<--	
2003US-475539P	P	20030602	<--	
2003WO-US38175	W	20031125		

CLASS

PATENT NO.	CLASS	PATENT FAMILY CLASSIFICATION CODES		
WO 2004047771	ICM	A61K		
	IPCI	A61K [ICM, 7]		
	IPCR	A61K0038-10 [I,A]; A61K0038-10 [I,C]; C07K0007-00 [I,C]; C07K0007-08 [I,A]		
	ECLA	C07K007/06A		
CA---2507045	IPCI	C07K0007-06 [ICM, 7]; C07K0007-00 [ICM, 7]; A61P0035-00 [ICS, 7]; A61K0038-05 [ICS, 7]; A61K0038-06 [ICS, 7]; C07K0005-06 [ICS, 7]; A61K0038-07 [ICS, 7]; A61K0038-08 [ICS, 7]; C07K0005-08 [ICS, 7]; C07K0005-10 [ICS, 7]; C07K0005-00 [ICS, 7]		
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	NCL	530/324.000		
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	IPCR	A61K0038-10 [I,A]; A61K0038-10 [I,C]; C07K0007-00 [I,C]; C07K0007-08 [I,A]		
	ECLA	C07K007/06A		
EP---1594521	IPCI	A61K0038-10 [ICM, 7]; A61K0038-08 [ICS, 7]; C07K0007-06 [ICS, 7]; C07K0007-08 [ICS, 7]; C07K0007-00 [ICS, 7]		
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CN---1741808	IPCI	A61K0038-10 [I,A]; A61K0038-08 [I,A]; C07K0007-06 [I,A]; C07K0007-08 [I,A]; C07K0007-00 [I,C]		
CN---1741809	IPCI	A61K0038-10 [I,A]; A61K0038-08 [I,A]; C07K0007-08 [I,A]; C07K0007-06 [I,A]; C07K0007-00 [I,C]		
JP2006514116	IPCI	C07K0005-06 [I,A]; A61K0038-00 [I,A]; A61P0035-00 [I,A]; A61K0045-00 [I,A]; C07K0005-097 [I,A]; C07K0005-117 [I,A]; C07K0005-00 [I,C]; C07K0007-06 [I,A]; C07K0007-00 [I,C]		
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NO2005003111	IPCI	C07K [ICM, 7]		
	ECLA	C07K007/06A		
OS MARPAT 141:33798				
AB	The invention discloses peptides which inhibit angiogenesis, cell migration, cell invasion and cell proliferation, as well as methods of making the peptides, pharmaceutical compns. containing the peptides, and methods of using the peptides and pharmaceutical compns. to treat diseases associated with aberrant vascularization, e.g. cancer.			
ST	peptide cell invasion migration proliferation inhibition; antitumor aberrant vascularization disease peptide prep			
IT	Sarcoma			

(cartilage chondrosarcoma; peptide inhibitors of angiogenesis, cell migration, cell invasion and cell proliferation, preparation, and compns. and therapeutic uses)

IT Cartilage, neoplasm
 (chondrosarcoma; peptide inhibitors of angiogenesis, cell migration, cell invasion and cell proliferation, preparation, and compns. and therapeutic uses)

IT Intestine, neoplasm
 (colon; peptide inhibitors of angiogenesis, cell migration, cell invasion and cell proliferation, preparation, and compns. and therapeutic uses)

IT Blood vessel
 (endothelium; peptide inhibitors of angiogenesis, cell migration, cell invasion and cell proliferation, preparation, and compns. and therapeutic uses)

IT Blood vessel, neoplasm
 Sarcoma
 (hemangiosarcoma; peptide inhibitors of angiogenesis, cell migration, cell invasion and cell proliferation, preparation, and compns. and therapeutic uses)

IT Angiogenesis
 Angiogenesis inhibitors
 Antitumor agents
 Brain, neoplasm
 Drug delivery systems
 Kidney, neoplasm
 Mammary gland, neoplasm
 Neoplasm
 Prostate gland, neoplasm
 (peptide inhibitors of angiogenesis, cell migration, cell invasion and cell proliferation, preparation, and compns. and therapeutic uses)

IT Endothelium
 (vascular; peptide inhibitors of angiogenesis, cell migration, cell invasion and cell proliferation, preparation, and compns. and therapeutic uses)

IT 701201-26-5D, biotinylated
 RL: BSU (Biological study, unclassified); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (peptide inhibitors of angiogenesis, cell migration, cell invasion and cell proliferation, preparation, and compns. and therapeutic uses)

IT 701200-82-0P 701201-01-6P
 RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)
 (peptide inhibitors of angiogenesis, cell migration, cell invasion and cell proliferation, preparation, and compns. and therapeutic uses)

IT 81658-55-1P 701200-81-9P 701200-83-1P
 701200-84-2P 701200-85-3P 701200-86-4P 701200-87-5P
 701200-88-6P 701200-89-7P 701200-90-0P
 701200-91-1P 701200-92-2P 701200-93-3P
 701200-94-4P 701200-95-5P 701200-96-6P 701200-97-7P 701200-98-8P
 701200-99-9P 701201-00-5P 701201-02-7P
 701201-03-8P 701201-04-9P 701201-05-0P
 701201-06-1P 701201-07-2P 701201-08-3P
 701201-09-4P 701201-10-7P 701201-11-8P
 701201-12-9P 701201-13-0P 701201-14-1P
 701201-15-2P 701201-16-3P 701201-17-4P
 701201-18-5P 701201-19-6P 701201-20-9P
 701201-21-0P 701201-22-1P 701201-23-2P 701201-24-3P
 701201-25-4P
 RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
 (peptide inhibitors of angiogenesis, cell migration, cell invasion and cell proliferation, preparation, and compns. and therapeutic uses)

IT 701201-28-7

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (peptide inhibitors of angiogenesis, cell migration, cell invasion and cell proliferation, preparation, and compns. and therapeutic uses)

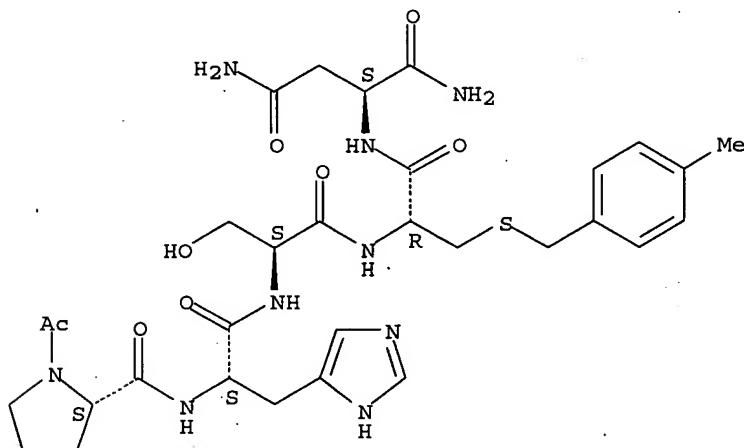
IT 98-88-4, Benzoyl chloride 100-39-0, Benzyl bromide 106-95-6, Allyl bromide, reactions 930-69-8 1212-08-4, S-Phenyl benzenethiosulfonate 2719-27-9, Cyclohexanoyl chloride 2937-50-0, Allyl chloroformate 2949-92-0, S-Methyl methanethiosulfonate 3282-30-2, Pivaloyl chloride 5271-67-0, 2-Thiophenecarbonyl chloride 6482-24-2, 2-Bromoethyl methylether 7031-27-8, (Phenylthio)acetyl chloride 10400-19-8, Nicotinoyl chloride 25644-88-6, S-Benzyl-L-cysteine sulfone 82911-69-1 262438-43-7 475150-36-8 701201-27-6
 RL: RCT (Reactant); RACT (Reactant or reagent)
 (peptide inhibitors of angiogenesis, cell migration, cell invasion and cell proliferation, preparation, and compns. and therapeutic uses)

IT 701200-82-0P
 RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)
 (peptide inhibitors of angiogenesis, cell migration, cell invasion and cell proliferation, preparation, and compns. and therapeutic uses)

RN 701200-82-0 HCPLUS

CN L-Aspartamide, 1-acetyl-L-prolyl-L-histidyl-L-seryl-S-[(4-methylphenyl)methyl]-L-cysteinyl - (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L26 ANSWER 3 OF 3 HCPLUS COPYRIGHT 2006 ACS on STN

AN 2003:243058 HCPLUS

DN 139:173332

ED Entered STN: 30 Mar 2003

TI Inhibition of integrin $\alpha 5\beta 1$ function with a small peptide (ATN-161) plus continuous 5-FU infusion reduces colorectal liver metastases and improves survival in mice

AU Stoeltzing, Oliver; Liu, Wenbiao; Reinmuth, Niels; Fan, Fan; Parry, Graham C.; Parikh, Alexander A.; McCarty, Marya F.; Bucana, Corazon D.; Mazar, Andrew P.; Ellis, Lee M.

CS Department of Cancer Biology, The University of Texas M.D. Anderson Cancer Center, Houston, TX, 77030-4009, USA

SO International Journal of Cancer (2003), 104(4), 496-503
 CODEN: IJCAW; ISSN: 0020-7136

PB Wiley-Liss, Inc.

DT Journal

LA English

CC 1-6 (Pharmacology)

- AB Integrin $\alpha 5\beta 1$ is expressed on activated endothelial cells and plays a critical role in tumor angiogenesis. We hypothesized that a novel integrin $\alpha 5\beta 1$ antagonist, ATN-161, would inhibit angiogenesis and growth of liver metastases in a murine model. We further hypothesized that combining ATN-161 with 5-fluorouracil (5-FU) chemotherapy would enhance the antineoplastic effect. Murine colon cancer cells (CT26) were injected into spleens of BALB/c mice to produce liver metastases. Four days thereafter, mice were given either ATN-161 (100 mg/kg, every 3rd day) or saline by i.p. injection, with or without combination of continuous-infusion 5-FU (100 mg/kg/2 wk), which was started on day 7. On day 20 after tumor cell inoculation, mice were killed and liver wts. and number of liver metastases were determined. A follow-up study on survival was also conducted in which mice were randomized to receive ATN-161, 5-FU or ATN-161+5-FU. Combination therapy with ATN-161+5-FU significantly reduced tumor burden (liver weight) and number of liver metastases ($p<0.02$). Liver tumors in the ATN-161 and ATN-161+5-FU groups had significantly fewer microvessels ($p<0.05$) than tumors in the control or 5-FU-treated groups. Unlike treatment with either agent alone, ATN-161+5-FU significantly increased tumor cell apoptosis and decreased tumor cell proliferation ($p<0.03$) and improved overall survival ($p<0.03$, log-rank test). Targeting integrin $\alpha 5\beta 1$ in combination with 5-FU infusion reduced liver metastases formation and improved survival in this colon cancer model. The enhancement of antineoplastic activity from the combination of anti-angiogenic therapy and chemotherapy may be a promising approach for treating metastatic colorectal cancer.
- ST ATN 161 fluorouracil colorectal cancer integrin
- IT Intestine, neoplasm
(colorectal; inhibition of integrin $\alpha 5\beta 1$ function with ATN-161 plus 5-FU infusion reduces colorectal liver metastases and improves survival in mice)
- IT Liver, neoplasm
(metastasis; inhibition of integrin $\alpha 5\beta 1$ function with ATN-161 plus 5-FU infusion reduces colorectal liver metastases and improves survival in mice)
- IT Drug interactions
(synergistic; inhibition of integrin $\alpha 5\beta 1$ function with ATN-161 plus 5-FU infusion reduces colorectal liver metastases and improves survival in mice)
- IT Integrins
RL: BSU (Biological study, unclassified); BIOL (Biological study)
($\alpha 5\beta 1$; inhibition of integrin $\alpha 5\beta 1$ function with ATN-161 plus 5-FU infusion reduces colorectal liver metastases and improves survival in mice)
- IT 262438-43-7, ATN 161
RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(inhibition of integrin $\alpha 5\beta 1$ function with ATN-161 plus 5-FU infusion reduces colorectal liver metastases and improves survival in mice)
- IT 51-21-8, 5-Fluorouracil
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(inhibition of integrin $\alpha 5\beta 1$ function with ATN-161 plus 5-FU infusion reduces colorectal liver metastases and improves survival in mice)
- RE.CNT 42 THERE ARE 42 CITED REFERENCES AVAILABLE FOR THIS RECORD
- RE
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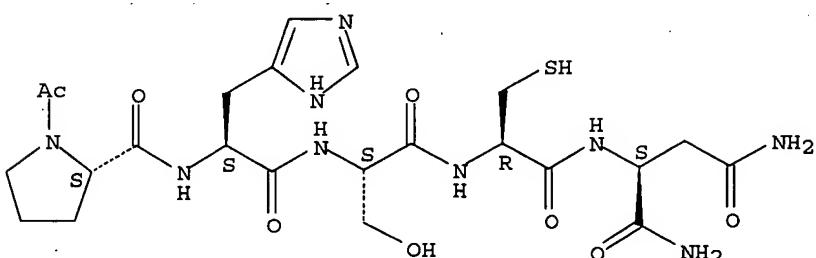
IT 262438-43-7, ATN 161

RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (inhibition of integrin $\alpha 5\beta 1$ function with ATN-161 plus 5-FU infusion reduces colorectal liver metastases and improves survival in mice)

RN 262438-43-7 HCAPLUS

CN L-Aspartamide, 1-acetyl-L-prolyl-L-histidyl-L-seryl-L-cysteinyl- (9CI)
 (CA INDEX NAME)

Absolute stereochemistry.



=> d all hitstr 127 tot

L27 ANSWER 1 OF 5 HCAPLUS COPYRIGHT 2006 ACS on STN

AN 2005:303191 HCAPLUS

DN 142:341966

ED Entered STN: 08 Apr 2005

TI Hydrogels used to deliver medicaments to the eye for the treatment of

posterior segment diseases
 IN Schultz, Clyde L.
 PA USA
 SO U.S. Pat. Appl. Publ., 7 pp., Cont.-in-part of U.S. Ser. No. 821,718.
 CODEN: USXXCO
 DT Patent
 LA English
 IC ICM A61K-0039/395
 ICS A61K-0038/24; A61K-0031/65; A61K-0009/14; A61K-0031/56; A61K-0031/075
 INCL 424486000; 514008000; 514012000; 424144100; 514182000; 514721000;
 514152000
 CC 63-6 (Pharmaceuticals)
 FAN.CNT 2

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US2005074497	A1	20050407	2004US-0971997	20041022
	US2005208102	A1	20050922	2004US-0821718	20040409
	US2005255144	A1	20051117	2005US-0102454	20050409
	WO2005110473	A2	20051124	2005WO-US12185	20050409
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG					
PRAI	2003US-461354P	P	20030409		
	2004US-0821718	A2	20040409		
	2004US-0971997	A2	20041022		

CLASS	PATENT NO.	CLASS	PATENT FAMILY CLASSIFICATION CODES
US 2005074497	US 2005074497	ICM	A61K-0039/395
		ICS	A61K-0038/24; A61K-0031/65; A61K-0009/14; A61K-0031/56; A61K-0031/075
		INCL	424486000; 514008000; 514012000; 424144100; 514182000; 514721000; 514152000
		IPCI	A61K0039-395 [ICM,7]; A61K0038-24 [ICS,7]; A61K0031-65 [ICS,7]; A61K0009-14 [ICS,7]; A61K0031-56 [ICS,7]; A61K0031-075 [ICS,7]
		IPCR	A61K0031-075 [I,A]; A61K0031-075 [I,C]; A61K0031-56 [I,A]; A61K0031-56 [I,C]; A61K0031-65 [I,A]; A61K0031-65 [I,C]
		NCL	424/486.000
		ECLA	A61K031/075; A61K031/56; A61K031/65
		IPCI	A61F0002-00 [ICM,7]; A61K0009-14 [ICS,7]
		IPCR	A61K0031-075 [I,A]; A61K0031-075 [I,C]; A61K0031-56 [I,A]; A61K0031-56 [I,C]; A61K0031-65 [I,A]; A61K0031-65 [I,C]
		NCL	424/427.000
US2005208102		ECLA	A61K031/075; A61K031/56; A61K031/65
		IPCI	A61F0002-00 [ICM,7]
		IPCR	A61K0031-075 [I,A]; A61K0031-075 [I,C]; A61K0031-56 [I,A]; A61K0031-56 [I,C]; A61K0031-65 [I,A]; A61K0031-65 [I,C]
		NCL	424/427.000
US2005255144		ECLA	A61K031/075; A61K031/56; A61K031/65
		IPCI	A61F0002-00 [ICM,7]
		IPCR	A61F0002-00 [I,A]; A61F0002-00 [I,C]
		NCL	424/428.000
WO2005110473	WO2005110473	IPCI	A61K0039-395 [ICM,7]; A61K0038-24 [ICS,7]; A61K0031-65 [ICS,7]; A61K0009-14 [ICS,7]; A61K0031-56 [ICS,7]; A61K0031-075 [ICS,7]

AB This invention provides a polymeric drug delivery system including a hydrogel containing one or more drugs for the treatment of a posterior segment disease. Exemplary drugs are anti-angiogenesis compds. for the treatment of macular degeneration. Allowing passive transference of this drug from

a dilute solution into the hydrogel produces the delivery system. The hydrogel, when placed in contact with the eye, delivers the drug. The delivery of the drug is sustained over an extended period of time, which is of particular utility in the eye, which is periodically flushed with tears. This sustained delivery accelerates the treatment process while avoiding potential damaging effects of localized delivery of high concns. of compds., e.g., from eye drops.

- ST hydrogel drug ophthalmic posterior segment delivery; angiogenesis inhibitor hydrogel eye macular degeneration treatment
- IT Thrombospondins
 - RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 - (1; hydrogels containing drugs for treatment of eye diseases in posterior segment)
- IT Platelet-derived growth factors
 - RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 - (BB; hydrogels containing drugs for treatment of eye diseases in posterior segment)
- IT Transcription factors
 - RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 - (HIF-1 α (hypoxia-inducible factor 1 α); hydrogels containing drugs for treatment of eye diseases in posterior segment)
- IT Polysaccharides, biological studies
 - RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 - (PS-K; hydrogels containing drugs for treatment of eye diseases in posterior segment)
- IT Vascular endothelial growth factor receptors
 - RL: BSU (Biological study, unclassified); BIOL (Biological study)
 - (VEGF trap; hydrogels containing drugs for treatment of eye diseases in posterior segment)
- IT Calreticulin
 - RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 - (amino-terminal fragment (vasostatin); hydrogels containing drugs for treatment of eye diseases in posterior segment)
- IT Growth factors, animal
 - Platelet-derived growth factors
 - RL: BSU (Biological study, unclassified); BIOL (Biological study)
 - (antagonists; hydrogels containing drugs for treatment of eye diseases in posterior segment)
- IT Epidermal growth factor receptors
 - RL: BSU (Biological study, unclassified); BIOL (Biological study)
 - (antibody; hydrogels containing drugs for treatment of eye diseases in posterior segment)
- IT Tea products
 - (green; hydrogels containing drugs for treatment of eye diseases in posterior segment)
- IT Angiogenesis inhibitors
 - Calcium channel blockers
 - Contact lenses
 - Cytotoxic agents
 - Human
 - Hydrogels
 - Sodium channel blockers
 - Viral vectors
 - Vision
 - (hydrogels containing drugs for treatment of eye diseases in posterior segment)
- IT CD59 (antigen)
- IT Fibronectins
- IT Hepatocyte growth factor
- IT Interleukin 12
- IT Interleukin 8

- Macrophage inflammatory protein 2 α
- Midkines
- Pleiotrophins
- Retinoids
- Tetracyclines
- Tumor necrosis factors
 - RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 - (hydrogels containing drugs for treatment of eye diseases in posterior segment)
- IT Polyesters, biological studies
 - RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 - (hydrogels containing drugs for treatment of eye diseases in posterior segment)
- IT Chemokines
 - RL: BSU (Biological study, unclassified); BIOL (Biological study)
 - (interferon γ -inducible protein-10, upregulators; hydrogels containing drugs for treatment of eye diseases in posterior segment)
- IT Proteins
 - RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 - (interferon-induced; hydrogels containing drugs for treatment of eye diseases in posterior segment)
- IT Eye, disease
 - (macula, degeneration, treatment of; hydrogels containing drugs for treatment of eye diseases in posterior segment)
- IT Antibodies and Immunoglobulins
 - RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 - (monoclonal, anti-VEGF Mab; hydrogels containing drugs for treatment of eye diseases in posterior segment)
- IT Antibodies and Immunoglobulins
 - RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 - (monoclonal, conjugates, with DOTA and radiolabeled; hydrogels containing drugs for treatment of eye diseases in posterior segment)
- IT Drug delivery systems
 - (ophthalmic; hydrogels containing drugs for treatment of eye diseases in posterior segment)
- IT Interferons
 - RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 - (α -2a; hydrogels containing drugs for treatment of eye diseases in posterior segment)
- IT Transforming growth factors
 - RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 - (α ; hydrogels containing drugs for treatment of eye diseases in posterior segment)
- IT Interferons
 - RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 - (α ; hydrogels containing drugs for treatment of eye diseases in posterior segment)
- IT Integrins
 - RL: BSU (Biological study, unclassified); BIOL (Biological study)
 - ($\alpha 5\beta 1$, antibody; hydrogels containing drugs for treatment of eye diseases in posterior segment)
- IT Transforming growth factors
 - RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 - (β ; hydrogels containing drugs for treatment of eye diseases in posterior segment)
- IT Interferons
 - RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(β ; hydrogels containing drugs for treatment of eye diseases in posterior segment)

IT Interferons
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (γ ; hydrogels containing drugs for treatment of eye diseases in posterior segment)

IT 99519-84-3, CAI
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (CAI; hydrogels containing drugs for treatment of eye diseases in posterior segment)

IT 212142-18-2, PTK 787
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (PTK 787; hydrogels containing drugs for treatment of eye diseases in posterior segment)

IT 92769-12-5, Proliferin
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (Proliferin; hydrogels containing drugs for treatment of eye diseases in posterior segment)

IT 106096-92-8, AFGF
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (aFGF; hydrogels containing drugs for treatment of eye diseases in posterior segment)

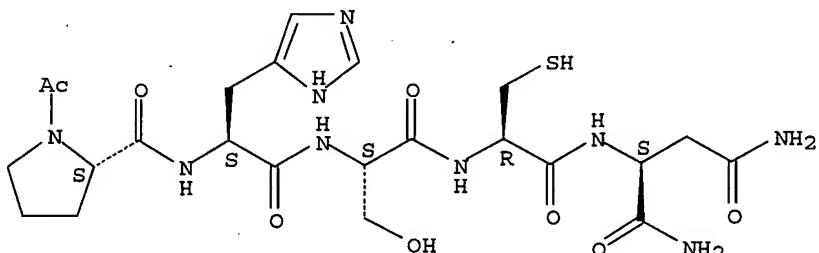
IT 127464-60-2, VEGF
 RL: BSU (Biological study, unclassified); BIOL (Biological study)
 (antibody; hydrogels containing drugs for treatment of eye diseases in posterior segment)

IT 50-35-1, Thalidomide 52-67-5, Penicillamine 362-07-2,
 2-Methoxyestradiol 446-72-0, Genistein 1861-40-1, Benefin 7559-04-8,
 Metarene 7753-60-8, Anecortave acetate 9000-94-6, Antithrombin
 9002-61-3, Choriogonadotropin 9002-62-4, Prolactin, biological studies
 9005-49-6D, Heparin, fragments, biological studies 9025-39-2, Heparinase
 9030-23-3, Plateletderived endothelial cell growth factor 10098-91-6D,
 Yttrium-90, mAbJ591-DOTA conjugates, labeled with, biological studies
 10540-29-1, Tamoxifen 15866-90-7, Col-3 16330-92-0 19171-19-8,
 Actimid 21416-67-1, Razoxane 24584-09-6, Dexrazoxane 33069-62-4,
 Paclitaxel 37270-94-3, Platelet factor-4 37300-21-3, Pentosan
 polysulfate 38101-59-6, IM862 58798-73-5, Tetrahydrocortisol sulfate
 59973-80-7, Aptosyn 60239-18-1D, DOTA, conjugates with mAbJ591,
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 Proliferin-related protein 105844-41-5, Plasminogen activator inhibitor
 106096-93-9, BFGF 112953-11-4, UCN-01 117570-53-3, DMXAA
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 growth factor 4 125313-92-0, Ro31-7453 129298-91-5, TNP-470
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 Progranulin 143011-72-7, Granulocyte colonystimulating factor
 144697-25-6 144941-41-3, Placental ribonuclease inhibitor 145266-99-5
 146426-40-6, Flavopiridol 148717-90-2, Squalamine 154039-60-8,
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 Medi-522 305838-77-1, Neovastat 324740-00-3, Vitaxin 339152-98-6,
 ABX-IL 8 347396-82-1, Ranibizumab 352423-07-5, Placental growth factor

365253-37-8, LY317615 402857-58-3, CEP-7055 443912-16-1, CC 7085
 443913-73-3, ZD6474 445467-72-1, ABT-510 557795-02-5, CP 564959
 564490-65-9 572921-97-2, Angiogenin 848784-95-2, AMG 001
 848784-98-5, MS 275291 848785-21-7, GBC 100 848785-35-3, ISV 120
 848785-37-5, Angioarrestin
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
 (Biological study); USES (Uses)
 (hydrogels containing drugs for treatment of eye diseases in posterior segment).

- IT 9003-07-0, Polypropylene 9004-34-6D, Cellulose, oxidized, regenerated
 26009-03-0, Polyglycolic acid 26023-30-3, Poly[oxy(1-methyl-2-oxo-1,2-ethanediyl)] 26124-68-5, Polyglycolic acid 26161-42-2 26680-10-4,
 Polylactide 26780-50-7, Polyglactin 910 29223-92-5, Poly-p-dioxanone
 30846-39-0, Poly(L-lactidecoglycolide) 31621-87-1, Polydioxanone
 33135-50-1, Poly(L-lactide) 35528-20-2, Vifilcon A 56551-60-1,
 Lidofilcon A 61463-79-4, Etafilcon A 114959-05-6, Poly-4-hydroxybutyrate 618109-09-4, Polymacon B 618109-14-1, Vasurfilcon A
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (hydrogels containing drugs for treatment of eye diseases in posterior segment)
- IT 141436-78-4, Protein kinase C
 RL: BSU (Biological study, unclassified); BIOL (Biological study)
 (inhibitors; hydrogels containing drugs for treatment of eye diseases in posterior segment)
- IT 262438-43-7, ATN-161
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
 (Biological study); USES (Uses)
 (hydrogels containing drugs for treatment of eye diseases in posterior segment)
- RN 262438-43-7 HCPLUS
 CN L-Aspartamide, 1-acetyl-L-prolyl-L-histidyl-L-seryl-L-cysteinyl- (9CI)
 (CA INDEX NAME)

Absolute stereochemistry.



- L27 ANSWER 2 OF 5 HCPLUS COPYRIGHT 2006 ACS on STN
 AN 2002:692596 HCPLUS
 DN 138:90056
 ED Entered STN: 13 Sep 2002
 TI Design and synthesis of the hydroxamic acid variants antitumorigenic and antimetastatic hydroxamate based Ac-PHSXX'-NH₂ sequences
 AU Sun, Yingchuan; Spatola, Arno F.
 CS Department of Chemistry and the Institute for Molecular Diversity and Drug Design, University of Louisville, Louisville, KY, 40292, USA
 SO Peptides: The Wave of the Future, Proceedings of the Second International and the Seventeenth American Peptide Symposium, San Diego, CA, United States, June 9-14, 2001 (2001), 799-800. Editor(s): Lebl, Michal; Houghten, Richard A. Publisher: American Peptide Society, San Diego, Calif.
 CODEN: 69DBAL; ISBN: 0-9715560-0-8
 DT Conference
 LA English
 CC 34-3 (Amino Acids, Peptides, and Proteins)

Section cross-reference(s): 1

AB A symposium report. Peptides MeCO-Pro-His-Ser-Xaa-Yaa-NH₂ [Xaa = Asp(NHOH), Glu(NHOH); Yaa = Leu, Asn], containing a hydroxamate group for superior metal-binding ability, were synthesized as potential antitumor and antimetastatic agents (biol. activity data not reported).

ST peptide hydroxamate prepn potential antitumor agent symposium

IT Antitumor agents

Neoplasm
(preparation of peptidyl hydroxamic acids as potential antitumor and antimetastatic agents)

IT Peptides, preparation
RL: SPN (Synthetic preparation); PREP (Preparation)
(preparation of peptidyl hydroxamic acids as potential antitumor and antimetastatic agents)

IT 483369-69-3P 483369-70-6P 483369-71-7P
RL: SPN (Synthetic preparation); PREP (Preparation)
(preparation of peptidyl hydroxamic acids as potential antitumor and antimetastatic agents)

RE.CNT 8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD

RE

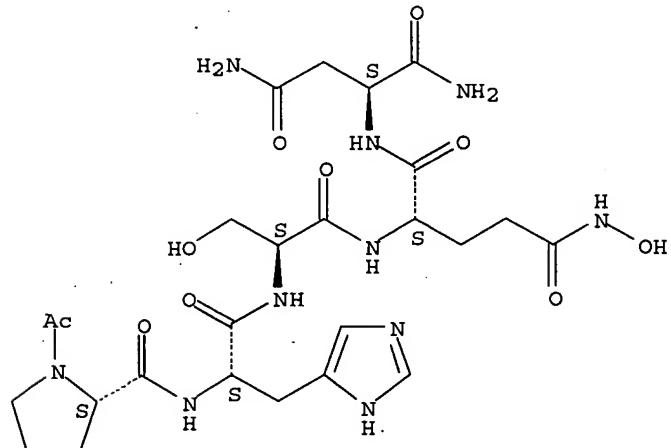
(1) Aota, S; J Biol Chem 1994, V269, P24756 HCPLUS
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(3) Livant, D; Cancer Res 2000, V60, P309 HCPLUS
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IT 483369-70-6P 483369-71-7P
RL: SPN (Synthetic preparation); PREP (Preparation)
(preparation of peptidyl hydroxamic acids as potential antitumor and antimetastatic agents)

RN 483369-70-6 HCPLUS

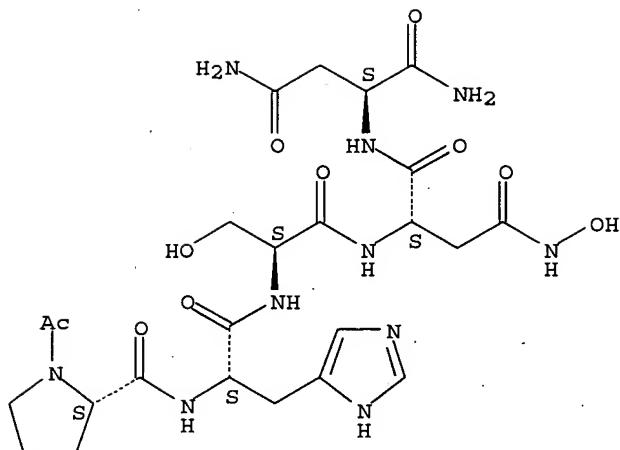
CN L-Aspartamide, 1-acetyl-L-prolyl-L-histidyl-L-seryl-N-hydroxy-L-glutaminyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



RN 483369-71-7 HCPLUS
CN L-Aspartamide, 1-acetyl-L-prolyl-L-histidyl-L-seryl-N-hydroxy-L-asparaginyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L27 ANSWER 3 OF 5 HCAPLUS COPYRIGHT 2006 ACS on STN

AN 2002:631705 HCAPLUS

DN 138:297158

ED Entered STN: 22 Aug 2002

TI Suppression of Tumor Recurrence and Metastasis by a Combination of the PHSCN Sequence and the Antiangiogenic Compound Tetrathiomolybdate in Prostate Carcinoma

AU van Golen, Kenneth L.; Bao, Liwei; Brewer, George J.; Pienta, Kenneth J.; Kamradt, Jeffrey M.; Livant, Donna L.; Merajver, Sofia D.

CS Division of Hematology and Oncology, Department of Internal Medicine, University of Michigan Comprehensive Cancer Center, Ann Arbor, MI, 48109-0948, USA

SO Neoplasia (New York, NY, United States) (2002), 4(5), 373-379
CODEN: NEOPFL; ISSN: 1522-8002

PB Nature Publishing Group

DT Journal

LA English

CC 1-6 (Pharmacology)

AB Plasma fibronectin-mediated invasion of human DU145 prostate cancer cell line was efficaciously inhibited in a rat tumor model by treatment with Ac-PHSCN-NH₂ peptide. Invasion of DU145 cells was stimulated by the PHSRN sequence of plasma fibronectin. However, PHSCN acts as a competitive inhibitor of PHSRN-mediated invasion. In the current study, we determined whether PHSCN could inhibit the recurrence and metastasis of DU145 tumors after excision of the primary tumor in an athymic nude mouse model. We demonstrated that mice treated thrice weekly with i.v. Ac-PHSCN-NH₂ peptide survived tumor-free for more than 30 wk post-primary tumor excision, whereas their untreated counterparts succumbed to recurrence and/or metastatic disease in significantly less time. Because of the universal requirement for angiogenesis in solid tumor growth, we tested the efficacy of copper deficiency induced by tetrathiomolybdate (TM) to retard tumor growth in the Dunning prostate cancer model. Significant reduction in size of the primary tumor was observed in mice rendered copper deficient. We sought to reduce tumor growth at the primary and metastatic sites by combining the anti-invasion Ac-PHSCN-NH₂ peptide with TM. Improved survival, fewer metastatic lesions, and excellent tolerability were observed with the combination therapy.

ST PHSCN peptide antiangiogenic tetrathiomolybdate prostate carcinoma recurrence metastasis

IT Prostate gland, neoplasm

(carcinoma, metastasis; suppression of tumor recurrence and metastasis by a combination of PHSCN sequence and the antiangiogenic compound tetrathiomolybdate in prostate carcinoma)

IT Prostate gland, neoplasm

(carcinoma; suppression of tumor recurrence and metastasis by a combination of PHSCN sequence and the antiangiogenic compound tetrathiomolybdate in prostate carcinoma)

IT Carcinoma
 (prostatic, metastasis; suppression of tumor recurrence and metastasis by a combination of PHSCN sequence and the antiangiogenic compound tetrathiomolybdate in prostate carcinoma)

IT Carcinoma
 (prostatic; suppression of tumor recurrence and metastasis by a combination of PHSCN sequence and the antiangiogenic compound tetrathiomolybdate in prostate carcinoma)

IT Angiogenesis inhibitors

Antitumor agents

Drug interactions

Human

(suppression of tumor recurrence and metastasis by a combination of PHSCN sequence and the antiangiogenic compound tetrathiomolybdate in prostate carcinoma)

IT 7440-50-8, Copper, biological studies

RL: BSU (Biological study, unclassified); BIOL (Biological study)
 (deficiency; efficacy of copper deficiency induced by tetrathiomolybdate (TM) to retard tumor growth in the Dunning prostate cancer model)

IT 16330-92-0 262438-43-7

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
 (Biological study); USES (Uses)
 (suppression of tumor recurrence and metastasis by a combination of PHSCN sequence and the antiangiogenic compound tetrathiomolybdate in prostate carcinoma)

RE.CNT 17 THERE ARE 17 CITED REFERENCES AVAILABLE FOR THIS RECORD

RE

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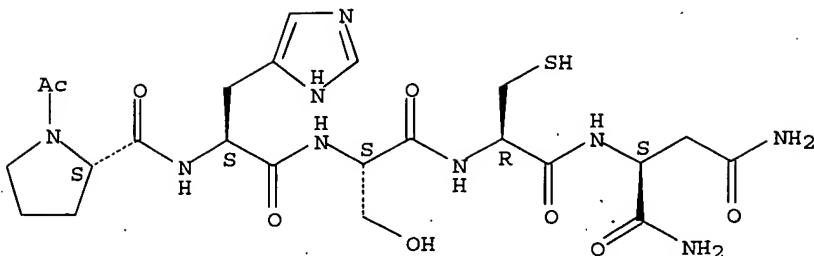
IT 262438-43-7

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
 (Biological study); USES (Uses)
 (suppression of tumor recurrence and metastasis by a combination of PHSCN sequence and the antiangiogenic compound tetrathiomolybdate in prostate carcinoma)

RN 262438-43-7 HCAPLUS

CN L-Aspartamide, 1-acetyl-L-prolyl-L-histidyl-L-seryl-L-cysteinyl- (9CI)
 (CA INDEX NAME)

Absolute stereochemistry.



L27 ANSWER 4 OF 5 HCAPLUS COPYRIGHT 2006 ACS on STN

AN 2002:555761 HCAPLUS

DN 137:121939

ED Entered STN: 26 Jul 2002

TI Compositions and methods for the use of fibronectin fragments in the diagnosis of cancer

IN Livant, Donna

PA The Regents of the University of Michigan, USA

SO PCT Int. Appl., 77 pp.

CODEN: PIXXD2

DT Patent

LA English

IC ICM G01N-0033/574

ICS A61K-0049/00; A61K-0051/00

CC 9-16 (Biochemical Methods)

Section cross-reference(s): 14

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO2002057786	A2	20020725	2002WO-US01189	20020115
	WO2002057786	A3	20031211		
	W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW			
	RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG			
	CA---2435320	AA	20020725	2002CA-2435320	20020115
	EP---1388013	A2	20040211	2002EP-0713418	20020115
	R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR			
PRAI	2001US-0765496	A	20010118		
	2002WO-US01189	W	20020115		

CLASS

	PATENT NO.	CLASS	PATENT FAMILY CLASSIFICATION CODES
WO	2002057786	ICM	G01N-0033/574
		ICS	A61K-0049/00; A61K-0051/00
		IPCI	G01N0033-574 [ICM, 7]; A61K0049-00 [ICS, 7]; A61K0051-00 [ICS, 7]
		IPCR	A61K0051-02 [I,C]; A61K0051-08 [I,A]; G01N0033-50 [I,A]; G01N0033-50 [I,C]; G01N0033-574 [I,A]; G01N0033-574 [I,C]
CA---	2435320	ECLA	A61K051/08; G01N033/50D4; G01N033/574V4
		IPCI	G01N0033-574 [ICM, 7]; A61K0049-00 [ICS, 7]; A61K0051-00 [ICS, 7]
EP---	1388013	IPCI	G01N0033-574 [ICM, 7]; A61K0049-00 [ICS, 7]; A61K0051-00 [ICS, 7]

IPCR A61K0051-02 [I,C]; A61K0051-08 [I,A]; G01N0033-50
[I,A]; G01N0033-50 [I,C]; G01N0033-574 [I,A];
G01N0033-574 [I,C]

- OS MARPAT 137:121939
- AB The present invention concerns the detection tumors *in vivo*, the imaging of tumors *in vivo*, and the imaging of cancerous tissue in pathol. samples. In particular the present invention incorporates the use of fibronectin fragments into these same detection and imaging methods.
- ST fibronectin cancer diagnosis imaging blood culture peptide radioisotope fluorescent
- IT Animal tissue
(autopsy sample; compns. and methods for use of fibronectin fragments in diagnosis of cancer)
- IT Radiography
(autoradiography; compns. and methods for use of fibronectin fragments in diagnosis of cancer)
- IT Animal tissue
Blood analysis
Blood plasma
Blood serum
Body fluid
Diagnosis
Epithelium
Fibroblast
Fluorescent substances
Human
Imaging
Mammary gland, neoplasm
Neoplasm
Pathology
Positron-emission tomography
Prostate gland
Rattus
Sample preparation
Scintigraphy
(compns. and methods for use of fibronectin fragments in diagnosis of cancer)
- IT Peptides, uses
Radionuclides, uses
RL: ARG (Analytical reagent use); PRP (Properties); ANST (Analytical study); USES (Uses)
(compns. and methods for use of fibronectin fragments in diagnosis of cancer)
- IT Fibronectins
RL: ANT (Analyte); BSU (Biological study, unclassified); DGN (Diagnostic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)
(degradation products; compns. and methods for use of fibronectin fragments in diagnosis of cancer)
- IT Drug delivery systems
(injections, i.v.; compns. and methods for use of fibronectin fragments in diagnosis of cancer)
- IT Skin
(keratinocyte; compns. and methods for use of fibronectin fragments in diagnosis of cancer)
- IT Culture media
(serum-free; compns. and methods for use of fibronectin fragments in diagnosis of cancer)
- IT Surgery
(tissue sample; compns. and methods for use of fibronectin fragments in diagnosis of cancer)
- IT 252229-84-8P
RL: ARG (Analytical reagent use); PRP (Properties); SPN (Synthetic preparation); ANST (Analytical study); PREP (Preparation); USES (Uses)
(compns. and methods for use of fibronectin fragments in diagnosis of cancer)
- IT 252230-05-0 262438-43-7 443305-20-2 443305-23-5

RL: ARG (Analytical reagent use); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)
 (compns. and methods for use of fibronectin fragments in diagnosis of cancer)

IT 252229-85-9P

RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
 (compns. and methods for use of fibronectin fragments in diagnosis of cancer)

IT 158622-13-0	252229-04-2	252229-05-3	252229-06-4	252229-07-5
252229-08-6	252229-09-7	252229-10-0	252229-11-1	252229-12-2
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252229-18-8	252229-19-9	252229-20-2	252229-21-3	252229-22-4
252229-23-5	252229-24-6	252229-25-7	252229-26-8	252229-27-9
252229-29-1	252229-30-4	252229-31-5	252229-32-6	252229-33-7
252229-34-8	252229-35-9	252229-36-0	252229-37-1	252229-38-2
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252229-71-3	252229-72-4	252229-73-5	252229-74-6	252229-75-7
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252229-81-5	252229-82-6	252229-83-7		

RL: PRP (Properties)

(unclaimed sequence; compns. and methods for the use of fibronectin fragments in the diagnosis of cancer)

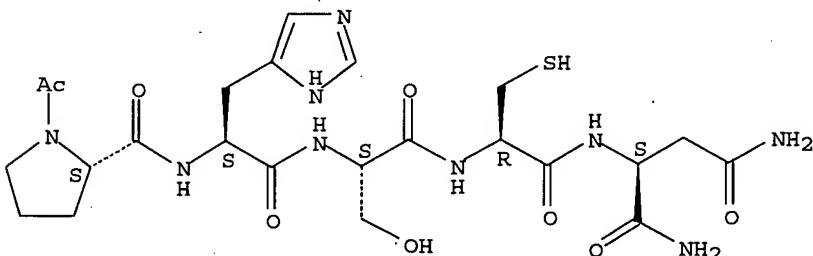
IT 262438-43-7 443305-23-5

RL: ARG (Analytical reagent use); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)
 (compns. and methods for use of fibronectin fragments in diagnosis of cancer)

RN 262438-43-7 HCPLUS

CN L-Aspartamide, 1-acetyl-L-prolyl-L-histidyl-L-seryl-L-cysteinyl- (9CI)
 (CA INDEX NAME)

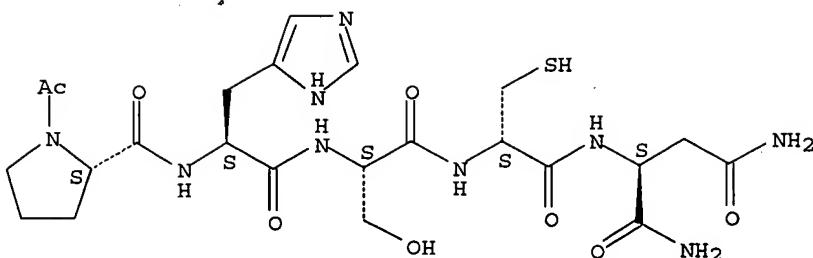
Absolute stereochemistry.



RN 443305-23-5 HCPLUS

CN L-Aspartamide, 1-acetyl-L-prolyl-L-histidyl-L-seryl-D-cysteinyl- (9CI)
 (CA INDEX NAME)

Absolute stereochemistry.



L27 ANSWER 5 OF 5 HCAPLUS COPYRIGHT 2006 ACS on STN
 AN 2000:102218 HCAPLUS
 DN 132:245978
 ED Entered STN: 13 Feb 2000
 TI Anti-invasive, antitumorigenic, and antimetastatic activities of the PHSCN sequence in prostate carcinoma
 AU Livant, Donna L.; Brabec, R. Kaye; Pienta, Kenneth J.; Allen, David L.; Kurachi, Kotoku; Markwart, Sonja; Upadhyaya, Ameet
 CS Department of Cell and Development Biology, University of Michigan Medical School, Ann Arbor, MI, 48109-0616, USA
 SO Cancer Research (2000), 60(2), 309-320
 CODEN: CNREA8; ISSN: 0008-5472
 PB AACR Subscription Office
 DT Journal
 LA English
 CC 1-6 (Pharmacology)
 AB Using naturally serum-free SU-ECM basement membranes as invasion substrates showed that plasma fibronectin was necessary to stimulate invasion by DU 145 human and metastatic MATLyLu (MLL) rat prostate carcinoma cells. This activity mapped to the PHSRN sequence, which induced invasion through $\alpha 5\beta 1$ integrin. PHSCN, a competitive inhibitor, blocked both PHSRN- and serum-induced invasion. Acetylated, amidated PHSCN (Ac-PHSCN-NH₂) was 30-fold more potent; however, Ac-HSPNC-NH₂ was inactive. Rats receiving injections s.c. with 100,000 MLL cells were treated systemically by i.v. injection three times weekly with 1 mg of either Ac-PHSCN-NH₂ or Ac-HSPNC-NH₂ beginning 24 h later, three times weekly with 1 mg of Ac-PHSCN-NH₂ beginning only after surgery to remove large (2 cm) MLL tumors, or were left untreated. MLL tumors grew rapidly in Ac-HSPNC-NH₂-treated and in untreated rats. MLL tumor growth in rats treated with Ac-PHSCN-NH₂ beginning 1 day after MLL cell injection was reduced by 99.9% during the first 16 days of treatment, although subsequent tumor growth occurred. MLL tumor cryosections immunostained with anti-PECAM-1 showed that Ac-PHSCN-NH₂ inhibited neovascularization by 12-fold during this time. Whether initiated after MLL cell injection or only after MLL tumor removal, Ac-PHSCN-NH₂ treatment reduced the nos. of MLL lung colonies and micrometastases by 40- to > 100-fold, whereas Ac-HSPNC-NH₂ was inactive. Thus, Ac-PHSCN-NH₂ may be a potent antitumorigenic and antimetastatic agent for postsurgical use prior to extensive metastasis.
 ST antitumorigenic antimetastatic PHSCN sequence prostate carcinoma; lung metastasis inhibitor prostate anticancer AcPHSCNNH2
 IT Angiogenesis inhibitors
 Protein sequences
 (anti-invasive, antitumorigenic, and antimetastatic activities of the PHSCN sequence in prostate carcinoma)
 IT Prostate gland
 Prostate gland
 (carcinoma, inhibitors; anti-invasive, antitumorigenic, and antimetastatic activities of the PHSCN sequence in prostate carcinoma)
 IT Lung, neoplasm
 (inhibitors, metastasis; anti-invasive, antitumorigenic, and antimetastatic activities of the PHSCN sequence in prostate carcinoma)

IT Antitumor agents
 (lung, metastasis; anti-invasive, antitumorigenic, and antimetastatic activities of the PHSCN sequence in prostate carcinoma)

IT Lung, neoplasm
 (metastasis, inhibitors; anti-invasive, antitumorigenic, and antimetastatic activities of the PHSCN sequence in prostate carcinoma)

IT Antitumor agents
 (prostate carcinoma; anti-invasive, antitumorigenic, and antimetastatic activities of the PHSCN sequence in prostate carcinoma)

IT 262438-43-7
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (anti-invasive, antitumorigenic, and antimetastatic activities of the PHSCN sequence in prostate carcinoma)

RE.CNT 58 THERE ARE 58 CITED REFERENCES AVAILABLE FOR THIS RECORD

RE

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IT 262438-43-7

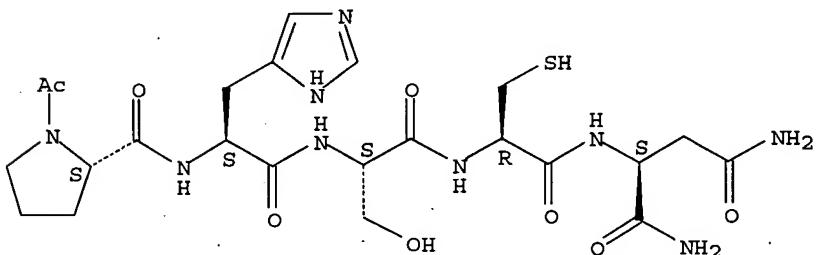
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(anti-invasive, antitumorigenic, and antimetastatic activities of the PHSCN sequence in prostate carcinoma)

RN 262438-43-7 HCPLUS

CN L-Aspartamide, 1-acetyl-L-prolyl-L-histidyl-L-seryl-L-cysteinyl- (9CI)
(CA INDEX NAME)

Absolute stereochemistry.



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FILE 'USPATFULL' ENTERED AT 07:02:45 ON 11 MAY 2006

CA INDEXING COPYRIGHT (C) 2006 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'USPAT2' ENTERED AT 07:02:45 ON 11 MAY 2006

CA INDEXING COPYRIGHT (C) 2006 AMERICAN CHEMICAL SOCIETY (ACS)

=> d bib abs hitstr l29 1-2

L29 ANSWER 1 OF 4 USPATFULL on STN

AN 2005:240102 USPATFULL

TI Hydrogels used to deliver medicaments to the eye for the treatment of posterior segment diseases

IN Schultz, Clyde L., Ponte Vedra, FL, UNITED STATES

PI US2005208102 A1 20050922

AI 2004US-0821718 A1 20040409 (10)

PRAI 2003US-461354P 20030409 (60)

DT Utility

FS APPLICATION

LREP FINCH IP LLC, P.O. BOX 1358, CONCORD, NH, 03302, US

CLMN Number of Claims: 20

ECL Exemplary Claim: 1

DRWN No Drawings

LN.CNT 502

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB This invention provides a polymeric drug delivery system including a hydrogel containing one or more drugs for the treatment of a posterior segment disease. Allowing passive transference of this drug from a

dilute solution into the hydrogel produces the delivery system. The hydrogel, when placed in contact with the eye, delivers the drug. The delivery of the drug is sustained over an extended period of time, which is of particular utility in the eye, which is periodically flushed with tears. This sustained delivery accelerates the treatment process while avoiding potential damaging effects of localized delivery of high concentrations of compounds, e.g., from eye drops.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

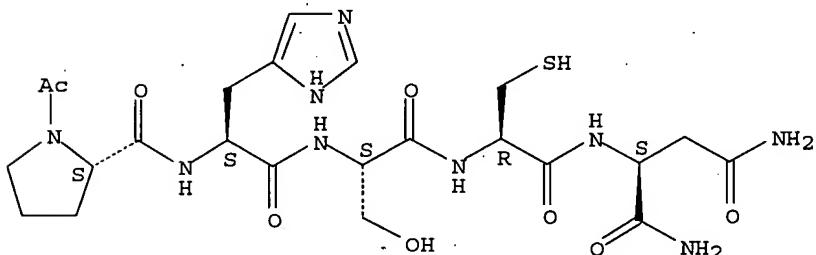
IT 262438-43-7, ATN-161

(hydrogels containing drugs for treatment of eye diseases in posterior segment)

RN 262438-43-7 USPATFULL

CN L-Aspartamide, 1-acetyl-L-proyl-L-histidyl-L-seryl-L-cysteinyl- (9CI)
(CA INDEX NAME)

Absolute stereochemistry.



L29 ANSWER 2 OF 4 USPATFULL on STN

AN 2005:87035 USPATFULL

TI Hydrogels used to deliver medicaments to the eye for the treatment of posterior segment diseases

IN Schultz, Clyde L., Ponte Vedra, FL, UNITED STATES

PI US2005074497 A1 20050407

AI 2004US-0971997 A1 20041022 (10)

RLI Continuation-in-part of Ser. No. 2004US-0821718, filed on 9 Apr 2004, PENDING

PRAI 2003US-461354P 20030409 (60)

DT Utility

FS APPLICATION

LREP CLARK & ELBING LLP, 101 FEDERAL STREET, BOSTON, MA, 02110

CLMN Number of Claims: 27

ECL Exemplary Claim: 1

DRWN No Drawings

LN.CNT 582

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB This invention provides a polymeric drug delivery system including a hydrogel containing one or more drugs for the treatment of a posterior segment disease. Exemplary drugs are anti-angiogenesis compounds for the treatment of macular degeneration. Allowing passive transference of this drug from a dilute solution into the hydrogel produces the delivery system. The hydrogel, when placed in contact with the eye, delivers the drug. The delivery of the drug is sustained over an extended period of time, which is of particular utility in the eye, which is periodically flushed with tears. This sustained delivery accelerates the treatment process while avoiding potential damaging effects of localized delivery of high concentrations of compounds, e.g., from eye drops.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

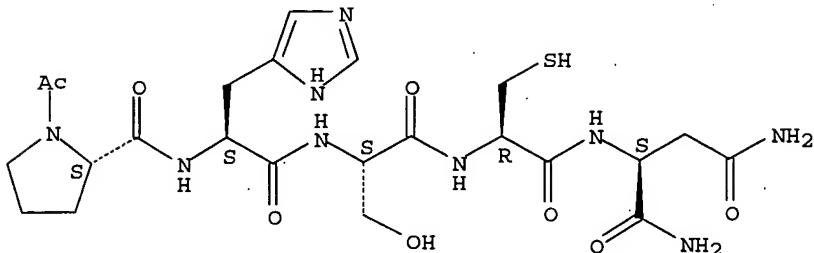
IT 262438-43-7, ATN-161

(hydrogels containing drugs for treatment of eye diseases in posterior segment)

RN 262438-43-7 USPATFULL

CN L-Aspartamide, 1-acetyl-L-prolyl-L-histidyl-L-seryl-L-cysteinyl- (9CI)
 (CA INDEX NAME)

Absolute stereochemistry.



=> d bib abs fhitstr 129 3-4

L29 ANSWER 3 OF 4 USPATFULL on STN

AN 2005:24260 USPATFULL

TI Peptides which target tumor and endothelial cells, compositions and uses thereof

IN Ternansky, Robert J., San Diego, CA, UNITED STATES
 Allan, Amy L., Encinitas, CA, UNITED STATES
 Gladstone, Patricia L., San Diego, CA, UNITED STATES
 Yoon, Won Hyung, San Diego, CA, UNITED STATES
 Parry, Graham, San Diego, CA, UNITED STATES
 Donate, Fernando, San Diego, CA, UNITED STATES
 Mazar, Andrew, San Diego, CA, UNITED STATES

PI US2005020810 A1 20050127

AI 2003US-0722843 A1 20031125 (10)

PRAI 2002US-429174P 20021125 (60)
 2003US-475539P 20030602 (60)

DT Utility

FS APPLICATION

LREP Sunil K. Singh, Dorsey & Whitney LLP, Intellectual Property Department,
 Four Embarcadero Center, Suite 3400, San Francisco, CA, 94111-4187

CLMN Number of Claims: 74

ECL Exemplary Claim: 1

DRWN No Drawings

LN.CNT 3884

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The present invention relates generally to peptide analogs of Ac--PHSCN--NH₂ which target tumor and endothelial cells and have anti-tumor, anti-angiogenic and anti-metastatic activity, methods of making these peptides, compositions thereof and methods of using these peptides and pharmaceutical compositions thereof to treat, prevent and detect diseases characterized by tumor growth, metastasis and angiogenesis. The peptide analogs may serve, inter alia, as carriers of radioactivity, PET-active compounds, toxins, fluorescent molecules and PEG molecules.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

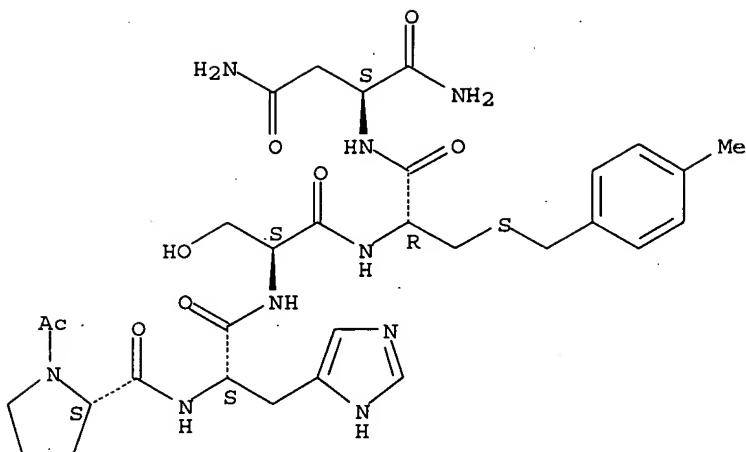
IT 701200-82-0P

(peptide inhibitors of angiogenesis, cell migration, cell invasion and cell proliferation, preparation, and compns. and therapeutic uses)

RN 701200-82-0 USPATFULL

CN L-Aspartamide, 1-acetyl-L-prolyl-L-histidyl-L-seryl-S-[(4-methylphenyl)methyl]-L-cysteinyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L29 ANSWER 4 OF 4 USPATFULL on STN

AN 2004:209805 USPATFULL

TI Peptides which inhibit angiogenesis, cell migration, cell invasion and cell proliferation, compositions and uses thereof

IN Allan, Amy L., Encinitas, CA, UNITED STATES

Donate, Fernando, San Diego, CA, UNITED STATES

Hopkins, Stephanie A., Poway, CA, UNITED STATES

Gladstone, Patricia L., San Diego, CA, UNITED STATES

Mazar, Andrew, San Diego, CA, UNITED STATES

O'Hare, Sean M., San Diego, CA, UNITED STATES

Parry, Graham, San Diego, CA, UNITED STATES

Plunkett, Marian, San Diego, CA, UNITED STATES

Ternansky, Robert J., San Diego, CA, UNITED STATES

Yoon, Won Hyung, San Diego, CA, UNITED STATES

PI US2004162239 A1 20040819

AI 2003US-0723144 A1 20031125 (10)

PRAI 2002US-429174P 20021125 (60)

2003US-475539P 20030602 (60)

DT Utility

FS APPLICATION

LREP COOLEY GODWARD, LLP, 3000 EL CAMINO REAL, 5 PALO ALTO SQUARE, PALO ALTO, CA, 94306

CLMN Number of Claims: 65

ECL Exemplary Claim: 1

DRWN 5 Drawing Page(s)

LN.CNT 3373

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The present invention relates generally to peptides, which inhibit angiogenesis, cell migration, cell invasion and cell proliferation, methods of making peptides, which inhibit angiogenesis, cell migration, cell invasion and cell proliferation, pharmaceutical compositions of these peptides and methods of using these peptides and pharmaceutical compositions of these peptides to treat diseases associated with aberrant vascularization.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

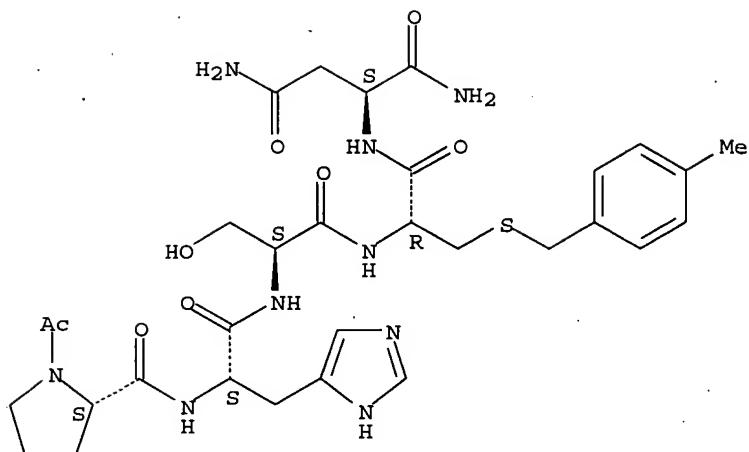
IT 701200-82-0P

(peptide inhibitors of angiogenesis, cell migration, cell invasion and cell proliferation, preparation, and compns. and therapeutic uses)

RN 701200-82-0 USPATFULL

CN L-Aspartamide, 1-acetyl-L-prolyl-L-histidyl-L-seryl-S-[(4-methylphenyl)methyl]-L-cysteinyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



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 L5 15 E3,E6
 E YOON WON/AU
 L6 10 E12
 E YOON WONHYING/AU
 E YOON WONHYUNG/AU
 E ALLAN A/AU
 L7 36 E3,E11
 E ALLAN AMY/AU
 L8 6 E5
 E GLADSTONE P/AU
 L9 28 E3-7
 E OHARE S/AU
 E O HARE S/AU
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 E O HARE SEAN/AU
 L11 8 E3-5
 E DONATE F/AU
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 E MAZAR A/AU
 L13 79 E3-4,E7-9
 E PARRY G/AU
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L29 4 L24

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